

## SEQUENCE LISTING

## (1) GENERAL INFORMATION:

- 5 (i) APPLICANT: Genentech, Inc.  
Garrard, Lisa J.  
Henner, Dennis J.  
Bass, Steven  
Greene, Ronald  
10 Lowman, Henry B.  
Wells, James A.  
Matthews, David J.
- (ii) TITLE OF INVENTION: Enrichment Method For Variant Proteins With  
15 Altered Binding Properties
- (iii) NUMBER OF SEQUENCES: 27
- (iv) CORRESPONDENCE ADDRESS:  
20 (A) ADDRESSEE: Genentech, Inc.  
(B) STREET: 460 Point San Bruno Blvd  
(C) CITY: South San Francisco  
(D) STATE: California  
(E) COUNTRY: USA  
25 (F) ZIP: 94080
- (v) COMPUTER READABLE FORM:  
(A) MEDIUM TYPE: 5.25 inch, 360 Kb floppy disk  
(B) COMPUTER: IBM PC compatible  
30 (C) OPERATING SYSTEM: PC-DOS/MS-DOS  
(D) SOFTWARE: patin (Genentech)
- (vi) CURRENT APPLICATION DATA:  
(A) APPLICATION NUMBER: 0  
35 (B) FILING DATE: 03-DEC-91  
(C) CLASSIFICATION:
- (vii) PRIOR APPLICATION DATA:  
(A) APPLICATION NUMBER: 07/743614  
40 (B) APPLICATION DATE: 09-Aug-91
- (vii) PRIOR APPLICATION DATA:  
(A) APPLICATION NUMBER: 07/715300  
(B) APPLICATION DATE: 14-June-91  
45
- (vii) PRIOR APPLICATION DATA:  
(A) APPLICATION NUMBER: 07/683400  
(B) APPLICATION DATE: 10-Apr-91  
50
- (vii) PRIOR APPLICATION DATA:  
(A) APPLICATION NUMBER: 07/621667  
(B) APPLICATION DATE: 03-Dec-1990  
55
- (viii) ATTORNEY/AGENT INFORMATION:  
(A) NAME: Benson, Robert H.  
(B) REGISTRATION NUMBER: 30,446  
(C) REFERENCE/DOCKET NUMBER: 645P4
- (ix) TELECOMMUNICATION INFORMATION:  
60 (A) TELEPHONE: 415/266-1489  
(B) TELEFAX: 415/952-9881  
(C) TELEX: 910/371-7168
- (2) INFORMATION FOR SEQ ID NO:1:
- 65 (i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 36 bases  
(B) TYPE: nucleic acid

- (C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:1:

5

GGCAGCTGTG GCTTCTAGAG TGGCGGCGGC TCTGGT 36

10 (2) INFORMATION FOR SEQ ID NO:2:

- (i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 36 bases  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO:2:

20

AGCTGTGGCT TCGGGCCCTT AGCATTTAAT GCGGTA 36

25 (2) INFORMATION FOR SEQ ID NO:3:

- (i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 33 bases  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

25

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO:3:

35

TTCAAAACG AAGGGCCCTT AATTAAAGCC AGA 33

(2) INFORMATION FOR SEQ ID NO:4:

- (i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 30 bases  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO:4:

CAATAATAAC GGGCTAGCCA AAAGAACTGG 30

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(2) INFORMATION FOR SEQ ID NO:5:

- (i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 24 bases  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO:5:

CACGACAGAA TTCCCGACTG GAAA 24

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(2) INFORMATION FOR SEQ ID NO:6:

- (i) SEQUENCE CHARACTERISTICS:

5 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:6:

10

15

20 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:7:

25

30

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:8:

40

(2) INFORMATION FOR SEQ ID NO:9:

45

50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:9:

55

60

65

ATGGACAAGG TGTCGACATA CCTGCGCATC GTG 33

5 (i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 36 bases  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

GGCAGCTGTG GCTTCTAGAG TGGCGGCGGC TCTGGT 36

(2) INFORMATION FOR SEQ ID NO:12:

25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:12:

30

(i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 12 amino acids  
(B) TYPE: amino acid  
(D) TOPOLOGY: linear

40 Gly Ser Cys Gly Phe Glu Ser Gly Gly Gly Ser Gly  
1 5 10 12

45 (i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 27 bases  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

CGGACTGGGC AGATATTCAA GCAGACC 27

(2) INFORMATION FOR SEQ ID NO:15:

65 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:15:

CTCAAGAACT ACGGGTTACC CTGACTGCTT CAGGAAGG 38

## (2) INFORMATION FOR SEQ ID NO:16:

- 5 (i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 30 bases  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

## 10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:16:

CGCATCGTGC AGTGCAGATC TGTGGAGGGC 30

## 15 (2) INFORMATION FOR SEQ ID NO:17:

- 20 (i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 66 bases  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

## 25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:17:

GTTACTCTAC TGCTTTCAGG AAGGACATGG ACNNSGTCNN SACANNSCTG 50  
30 NNSATCGTGC AGTGCA 66

## 35 (2) INFORMATION FOR SEQ ID NO:18:

- (i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 64 bases  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

## 40 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:18:

45 GATCTGCACT GCACGATSNN CAGSNNTGTS NNGACSNNGT CCATGTCCTT 50  
CCTGAAGCAG TAGA 64

## 50 (2) INFORMATION FOR SEQ ID NO:19:

- 55 (i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 25 bases  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

## 60 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:19:

GCCTTTGACA GGTACCAGGA GTTTG 25

## 65 (2) INFORMATION FOR SEQ ID NO:20:

- (i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 33 bases

5 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:20:

10 (2) INFORMATION FOR SEQ ID NO:21:

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:21:

25 GGCCTTTGAC ACGTAC/66

(2) INFORMATION FOR SEQ ID NO:22:

35 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:22:

40

SNNGAGCC 58

45 (2) INFORMATION FOR SEQ ID NO:23:

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:23:

GCNNSGTGCA GTGCA 65

(2) INFORMATION FOR SEQ ID NO:24:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 64 bases
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:24:

5 GATCTGCACT GCACSNNGCG CAGGTASNNG CTGACCTTSN NCATGTCCTT 50  
SNNGAAGCAG TAGA 64

10 (2) INFORMATION FOR SEQ ID NO:25:

(i) SEQUENCE CHARACTERISTICS:

15 (A) LENGTH: 2178 bases  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:25:

20 ATGAAAAAGA ATATCGCATT TCTTCTTGCA TCTATGTTTCG TTTTCTCTAT 50  
TGCTACAAAC GCGTACGCTG ATATCCAGAT GACCCAGTCC CCGAGCTCCC 100  
25 TGTCCGCCTC TGTGGGCGAT AGGGTCACCA TCACCTGCCG TGCCAGTCAG 150  
GATGTGAATA CTGCTGTAGC CTGGTATCAA CAGAAACCAG GAAAAGCTCC 200  
30 GAAACTACTG ATTTACTCGG CATCCTTCCT CTACTCTGGA GTCCCTTCTC 250  
35 GCTTCTCTGG ATCCAGATCT GGGACGGATT TCACTCTGAC CATCAGCAGT 300  
CTGCAGCCGG AAGACTTCGC AACTTATTAC TGTCAGCAAC ATTATACTAC 350  
40 TCCTCCCACG TTCGGACAGG GTACCAAGGT GGAGATCAAA CGAACTGTGG 400  
CTGCACCATC TGTCTTCATC TTCCCGCCAT CTGATGAGCA GTTGAAATCT 450  
45 GGAAGTGCCT CTGTTGTGTG CCTGCTGAAT AACTTCTATC CGAGAGAGGC 500  
CAAAGTACAG TGGAAGGTGG ATAACGCCCT CCAATCGGGT AACTOCCAGG 550  
AGAGTGTCAC AGAGCAGGAC AGCAAGGACA GCACCTACAG CCTCAGCAGC 600  
55 ACCCTGACGC TGAGCAAAGC AGACTACGAG AAACACAAAG TCTACGCCTG 650  
CGAAGTCACC CATCAGGGCC TGAGCTCGCC CGTCACAAAG AGCTTCAACA 700  
60 GGGGAGAGTG TTAAGCTGAT CCTCTACGCC GGACGCATCG TGGCCCTAGT 750  
65 ACGCAAGTTC ACGTAAAAAG GGTATCTAGA GGTGAGGTG ATTTTATGAA 800

AAAGAATATC GCATTTCTTC TTGCATCTAT GTTCGTTTTT TCTATTGCTA 850  
5 CAAACGCGTA CGCTGAGGTT CAGCTGGTGG AGTCTGGCGG TGGCCTGGTG 900  
CAGCCAGGGG GCTCACTCCG TTTGTCCTGT GCAGCTTCTG GCTTCAACAT 950  
10 TAAAGACACC TATATACACT GGGTGCGTCA GGCCCCGGT AAGGGCCTGG 1000  
AATGGGTTGC AAGGATTTAT CCTACGAATG GTTATACTAG ATATGCCGAT 1050  
15 AGCGTCAAGG GCCCTTTCAC TATAAGCGCA GACACATCCA AAAACACAGC 1100  
CTACCTGCAG ATGAACAGCC TGCCTGCTGA GGACACTGCC GTCTATTATT 1150  
20 GTTCTAGATG GGGAGGGGAC GGCTTCTATG CTATGGACTA CTGGGGTCAA 1200  
GGAACCCTGG TCACCGTCTC CTCGGCCTCC ACCAAGGGCC CATCGGTCTT 1250  
25 CCCCCTGGCA CCCTCCTCCA AGAGCACCTC TGGGGGCACA GCGGCCCTGG 1300  
30 GCTGCCTGGT CAAGGACTAC TTCCCCGAAC CGGTGACGGT GTCGTGGAAC 1350  
TCAGGCGCCC TGACCAGCGG CGTGACACCC TTCCCGGCTG TCCTACAGTC 1400  
35 CTCAGGACTC TACTCCCTCA GCAGCGTGGT GACTGTGCCC TCTAGCAGCT 1450  
TGGGCACCCA GACCTACATC TGCAACGTGA ATCACAAGCC CAGCAACACC 1500  
40 AAGGTGGACA AGAAAGTTGA GCCCAAATCT TGTGACAAA CTCACACAGG 1550  
45 GCCCTTCGTT TGTGAATATC AAGGCCAATC GTCTGACCTG CCTCAACCTC 1600  
CTGTCAATGC TGGCGGCGGC TCTGGTGGTG GTTCTGGTGG CGGCTCTGAG 1650  
50 GGTGGTGGCT CTGAGGGTGG CGGTTCTGAG GGTGGCGGCT CTGAGGGAGG 1700  
CGGTTCCGGT GGTGGCTCTG GTTCCGGTGA TTTTGATTAT GAAAAGATGG 1750  
55 CAAACGCTAA TAAGGGGGCT ATGACCGAAA ATGCCGATGA AAACGCGCTA 1800  
60 CAGTCTGACG CTAAAGGCAA ACTTGATTCT GTCGCTACTG ATTACGGTGC 1850  
TGCTATCGAT GGTTTCATTG GTGACGTTTC CGGCCTTGCT AATGGTAATG 1900  
65 GTGCTACTGG TGATTTTGCT GGCTCTAATT CCCAAATGGC TCAAGTCGGT 1950



5 GACGGTGATA ATTCACCTTT AATGAATAAT TTCCGTCAAT ATTTACCTTC 2000  
 CCTCCCTCAA TCGGTTGAAT GTCGCCCTTT TGTCTTTAGC GCTGGTAAAC 2050  
 CATATGAATT TTCTATTGAT TGTGACAAAA TAAACTTATT CCGTGGTGTC 2100  
 10 TTTGCGTTTC TTTTATATGT TGCCACCTTT ATGTATGTAT TTTCTACGTT 2150  
 TGCTAACATA CTGCGTAATA AGGAGTCT 2178

(2) INFORMATION FOR SEQ ID NO:26:

- (i) SEQUENCE CHARACTERISTICS:  
 (A) LENGTH: 237 amino acids  
 (B) TYPE: amino acid  
 (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:26:

25 Met Lys Lys Asn Ile Ala Phe Leu Leu Ala Ser Met Phe Val Phe  
 1 5 10 15  
 Ser Ile Ala Thr Asn Ala Tyr Ala Asp Ile Gln Met Thr Gln Ser  
 20 25 30  
 30 Pro Ser Ser Leu Ser Ala Ser Val Gly Asp Arg Val Thr Ile Thr  
 35 40 45  
 Cys Arg Ala Ser Gln Asp Val Asn Thr Ala Val Ala Trp Tyr Gln  
 50 55 60  
 35 Gln Lys Pro Gly Lys Ala Pro Lys Leu Leu Ile Tyr Ser Ala Ser  
 65 70 75  
 40 Phe Leu Tyr Ser Gly Val Pro Ser Arg Phe Ser Gly Ser Arg Ser  
 80 85 90  
 Gly Thr Asp Phe Thr Leu Thr Ile Ser Ser Leu Gln Pro Glu Asp  
 95 100 105  
 45 Phe Ala Thr Tyr Tyr Cys Gln Gln His Tyr Thr Thr Pro Pro Thr  
 110 115 120  
 Phe Gly Gln Gly Thr Lys Val Glu Ile Lys Arg Thr Val Ala Ala  
 125 130 135  
 50 Pro Ser Val Phe Ile Phe Pro Pro Ser Asp Glu Gln Leu Lys Ser  
 140 145 150  
 55 Gly Thr Ala Ser Val Val Cys Leu Leu Asn Asn Phe Tyr Pro Arg  
 155 160 165  
 Glu Ala Lys Val Gln Trp Lys Val Asp Asn Ala Leu Gln Ser Gly  
 170 175 180  
 60 Asn Ser Gln Glu Ser Val Thr Glu Gln Asp Ser Lys Asp Ser Thr  
 185 190 195  
 Tyr Ser Leu Ser Ser Thr Leu Thr Leu Ser Lys Ala Asp Tyr Glu  
 200 205 210  
 65 Lys His Lys Val Tyr Ala Cys Glu Val Thr His Gln Gly Leu Ser  
 215 220 225

004643460

(2) INFORMATION FOR SEQ ID NO:27:

(i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 461 amino acids  
(B) TYPE: amino acid  
(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:27:

15	Met	Lys	Lys	Asn	Ile	Ala	Phe	Leu	Leu	Ala	Ser	Met	Phe	Val	Phe	15
	1				5					10						
	Ser	Ile	Ala	Thr	Asn	Ala	Tyr	Ala	Glu	Val	Gln	Leu	Val	Glu	Ser	30
					20					25						
20	Gly	Gly	Gly	Leu	Val	Gln	Pro	Gly	Gly	Ser	Leu	Arg	Leu	Ser	Cys	45
					35					40						
	Ala	Ala	Ser	Gly	Phe	Asn	Ile	Lys	Asp	Thr	Tyr	Ile	His	Trp	Val	60
					50					55						
25	Arg	Gln	Ala	Pro	Gly	Lys	Gly	Leu	Glu	Trp	Val	Ala	Arg	Ile	Tyr	75
					65					70						
	Pro	Thr	Asn	Gly	Tyr	Thr	Arg	Tyr	Ala	Asp	Ser	Val	Lys	Gly	Arg	90
					80					85						
30	Phe	Thr	Ile	Ser	Ala	Asp	Thr	Ser	Lys	Asn	Thr	Ala	Tyr	Leu	Gln	105
					95					100						
35	Met	Asn	Ser	Leu	Arg	Ala	Glu	Asp	Thr	Ala	Val	Tyr	Tyr	Cys	Ser	120
					110					115						
	Arg	Trp	Gly	Gly	Asp	Gly	Phe	Tyr	Ala	Met	Asp	Tyr	Trp	Gly	Gln	135
					125					130						
40	Gly	Thr	Leu	Val	Thr	Val	Ser	Ser	Ala	Ser	Thr	Lys	Gly	Pro	Ser	150
					140					145						
	Val	Phe	Pro	Leu	Ala	Pro	Ser	Ser	Lys	Ser	Thr	Ser	Gly	Gly	Thr	165
					155					160						
45	Ala	Ala	Leu	Gly	Cys	Leu	Val	Lys	Asp	Tyr	Phe	Pro	Glu	Pro	Val	180
					170					175						
	Thr	Val	Ser	Trp	Asn	Ser	Gly	Ala	Leu	Thr	Ser	Gly	Val	His	Thr	195
					185					190						
50	Phe	Pro	Ala	Val	Leu	Gln	Ser	Ser	Gly	Leu	Tyr	Ser	Leu	Ser	Ser	210
					200					205						
55	Val	Val	Thr	Val	Pro	Ser	Ser	Ser	Leu	Gly	Thr	Gln	Thr	Tyr	Ile	225
					215					220						
	Cys	Asn	Val	Asn	His	Lys	Pro	Ser	Asn	Thr	Lys	Val	Asp	Lys	Lys	240
					230					235						
60	Val	Glu	Pro	Lys	Ser	Cys	Asp	Lys	Thr	His	Thr	Gly	Pro	Phe	Val	255
					245					250						
65	Cys	Glu	Tyr	Gln	Gly	Gln	Ser	Ser	Asp	Leu	Pro	Gln	Pro	Pro	Val	270
					260	</										

Gly Gly Gly Ser Glu Gly Gly Gly Ser Glu Gly Gly Gly Ser Glu  
 290 295 300  
 5 Gly Gly Gly Ser Gly Gly Gly Ser Gly Ser Gly Asp Phe Asp Tyr  
 305 310 315  
 Glu Lys Met Ala Asn Ala Asn Lys Gly Ala Met Thr Glu Asn Ala  
 320 325 330  
 10 Asp Glu Asn Ala Leu Gln Ser Asp Ala Lys Gly Lys Leu Asp Ser  
 335 340 345  
 Val Ala Thr Asp Tyr Gly Ala Ala Ile Asp Gly Phe Ile Gly Asp  
 350 355 360  
 15 Val Ser Gly Leu Ala Asn Gly Asn Gly Ala Thr Gly Asp Phe Ala  
 365 370 375  
 Gly Ser Asn Ser Gln Met Ala Gln Val Gly Asp Gly Asp Asn Ser  
 380 385 390  
 20 Pro Leu Met Asn Asn Phe Arg Gln Tyr Leu Pro Ser Leu Pro Gln  
 395 400 405  
 25 Ser Val Glu Cys Arg Pro Phe Val Phe Ser Ala Gly Lys Pro Tyr  
 410 415 420  
 Glu Phe Ser Ile Asp Cys Asp Lys Ile Asn Leu Phe Arg Gly Val  
 425 430 435  
 30 Phe Ala Phe Leu Leu Tyr Val Ala Thr Phe Met Tyr Val Phe Ser  
 440 445 450  
 35 Thr Phe Ala Asn Ile Leu Arg Asn Lys Glu Ser  
 455 460 461

600  
 400  
 200  
 0